```
  1 GGCACGAGGAGATCTAGGTTCAAATTAATGTTGCCCCTAGTGGTAAAGGACAGAGACCCTCAGACTGATGAAATGCGCTCAGAATTACTT
 91 AGACAAAGCGGATATTTGCCACTCTCTTCCCCTTTTCCTGTGTTTTTGTAGTGAAGAGACCTGAAAGAAAAAAGTAGGGAGAACATAATG
                                                    *  *

181 AGAACAAATACGGTAATCTCTTCATTTGCTAGTTCAAGTGCTGGACTTGGGACTTAGGAGGGGCAATGGAGCCGCTTAGTGCCTACATCT
                                                          *

271 GACTTGGACTGAAATATAGGTGAGAGACAAGATTGTCTCATATCCGGGGAAATCATAACCTATGACTAGGACGGGAAGAGGAAGCACTGC
           *                                                      *

361 CTTTACTTCAGTGGGAATCTCGGCCTCAGCCTGCAAGCCAAGTGTTCACAGTGAGAAAAGCAAGAGAATAAGCTAATACTCCTGTCCTGA
                                                    *                                       *

451 ACAAGGCAGCGGCTCCTTGGTAAAGCTACTCCTTGATCGATCCTTTGCACCGGATTGTTCAAAGTGGACCCCAGGGGAGAAGTCGGAGCA
                                          *

541 AAGAACTTACCACCAAGCAGTCCAAGAGGCCCAGAAGCAAACCTGGAGGTGAGACCCAAAGAAAGCTGGAACCATGCTGACTTTGTACAC
                                                L  E  V  R  P  K  E  S  W  N  H  A  D  F  V  H     16

631 TGTGAGGACACAGAGTCTGTTCCTGGAAAGCCCAGTGTCAACGCAGATGAGGAAGTCGGAGGTCCcCAAATCTGCCGTGTATGTGGGGAC
    C  E  D  T  E  S  V  P  G  K  P  S  V  N  A  D  E  E  V  G  G  P  Q  I  C  R  V  C  G  D     46

721 AAGGCCACTGGCTATCACTTCAATGTCATGACATGTGAAGGATGCAAGGGCtTTTTCAGGAGGGCCATGAAACGCAACGCCCGGCTGAGG
    K  A  T  G  Y  H  F  N  V  M  T  C  E  G  C  K  G  F  F  R  R  A  M  K  R  N  A  R  L  R     76

811 TGCCCcTTCCGGAAGGGCGCCTGCGAGATCACCCGGAAGACCCGGCGACAGTGCCAGGCCTGCCGCCTGCGCAAGTGCCTGGAGAGCGGC
    C  P  F  R  K  G  A  C  E  I  T  R  K  T  R  R  Q  C  Q  A  C  R  L  R  K  C  L  E  S  G    106

901 ATGAAGAAGGAGATGATCATGTCCGACGAGGCCGTGGAGGAGAGGCGGGCCTTGATCAAGCGGAAGAAAAGTGAACGGACAGGGACTCAG
    M  K  K  E  M  I  M  S  D  E  A  V  E  E  R  R  A  L  I  K  R  K  K  S  E  R  T  G  T  Q    136
```

FIGURE 1A-1

```
991 CCACTGGGAGTGCAGGGGCTGACAGAGGAGCAGCGGATGATGATCAGGGAGCTGATGGACGCTCAGATGAAAACCTTTGACACTACCTTC
    P  L  G  V  Q  G  L  T  E  E  Q  R  M  M  I  R  E  L  M  D  A  Q  M  K  T  F  D  T  T  F    166

1081 TCCCATTTCAAGAATTTCCGGCTGCCAGGGGTGCTTAGCAGTGGCTGCGAGTTGCCAGAGCCTCTGCAGGCCCCATCGAGGGAAGAAGCT
     S  H  F  K  N  F  R  L  P  G  V  L  S  S  G  C  E  L  P  E  P  L  Q  A  P  S  R  E  E  A    196

1171 GCCAAGTGGAGCCAGGTCCGGAAAGATCTGTGCTCTTTGAAGGTCTCTCTGCAAGCTGCGGGGGGAGGATGGCAGTGTCTGGAACTACAA
     A  K  W  S  Q  V  R  K  D  L  C  S  L  K  V  S  L  Q  A  A  G  G  G  W  Q  C  L  E  L  Q    226

1261 ACNCCCAGCCGACAGTGGCGGAAAGAGATCTTCTCCCTGCTGCCCCACATGGCTGACATGTCAACCTACATGTTCAAAGGCATCATCAGC
     T  P  S  R  Q  W  R  K  E  I  F  S  L  L  P  H  M  A  D  M  S  T  Y  M  F  K  G  I  I  S    256

1351 TTTGCCAAAGTCATCTCCTACTTCAGGGACTTGCCCATCGAGGACCAGATCTCCCTGCTGAAGGGGGCCGCTTTCGAGCTGTGTCAACTG
     F  A  K  V  I  S  Y  F  R  D  L  P  I  E  D  Q  I  S  L  L  K  G  A  A  F  E  L  C  Q  L    286

1441 AGATTCAACACAGTGTTCAACGCGGAGACTGGAACCTGGGAGTGTGGCCGGCTGTCCTACTGCTTGGAAGACACTGCAGGTGGCTTCCAG
     R  F  N  T  V  F  N  A  E  T  G  T  W  E  C  G  R  L  S  Y  C  L  E  D  T  A  G  G  F  Q    316

1531 CAACTTCTACTGGAGCCCATGCTGAAATTCCACTACATGCTGAAGAAGCTGCAGCTGCATGAGGAGGAGTATGTGCTGATGCAGGCCATC
     Q  L  L  L  E  P  M  L  K  F  H  Y  M  L  K  K  L  Q  L  H  E  E  E  Y  V  L  M  Q  A  I    346

1621 TCCCTCTTCTCCCCAGACCGCCCAGGTGTGCTGCAGCACCGCGTGGTGGACCAGCTGCAGGAGCAATTCGCCATTACTCTGAAGTCCTAC
     S  L  F  S  P  D  R  P  G  V  L  Q  H  R  V  V  D  Q  L  Q  E  Q  F  A  I  T  L  K  S  Y    376

1711 ATTGAATGCAATCGGCCCCAGCCTGCTCATAGGTTCTTGTTCCTGAAGATCATGGCTATGCTCACCGAGCTCCGCAGCATCAATGCTCAG
     I  E  C  N  R  P  Q  P  A  H  R  F  L  F  L  K  I  M  A  M  L  T  E  L  R  S  I  N  A  Q    406

1801 CACACCCAGCGGCTGCTGCGCATCCAGGACATACACCCCTTTGCTACGCCCCTCATGCAGGAGTTGTTCGGCATCACAGGTAGCTGAGCG
     H  T  Q  R  L  L  R  I  Q  D  I  H  P  F  A  T  P  L  M  Q  E  L  F  G  I  T  G  S  *      434

1891 GCTGCCTTGGGTGACACCTTCGAGAGGCAGCCAGACCCAGAGCCCTCTGAGCCGGCACTCCCGGGCCAAGACAGATGGACACTGCCAAGA
1981 GCCGACAATGCCCTGCTGGCCTGTCTCCCTAGGGAATTCCTGCTATGACAGCTGGCTAGCATTCCTCAGGAAGGACATGGGGTGCCCC  2068
```

FIGURE 1A-2

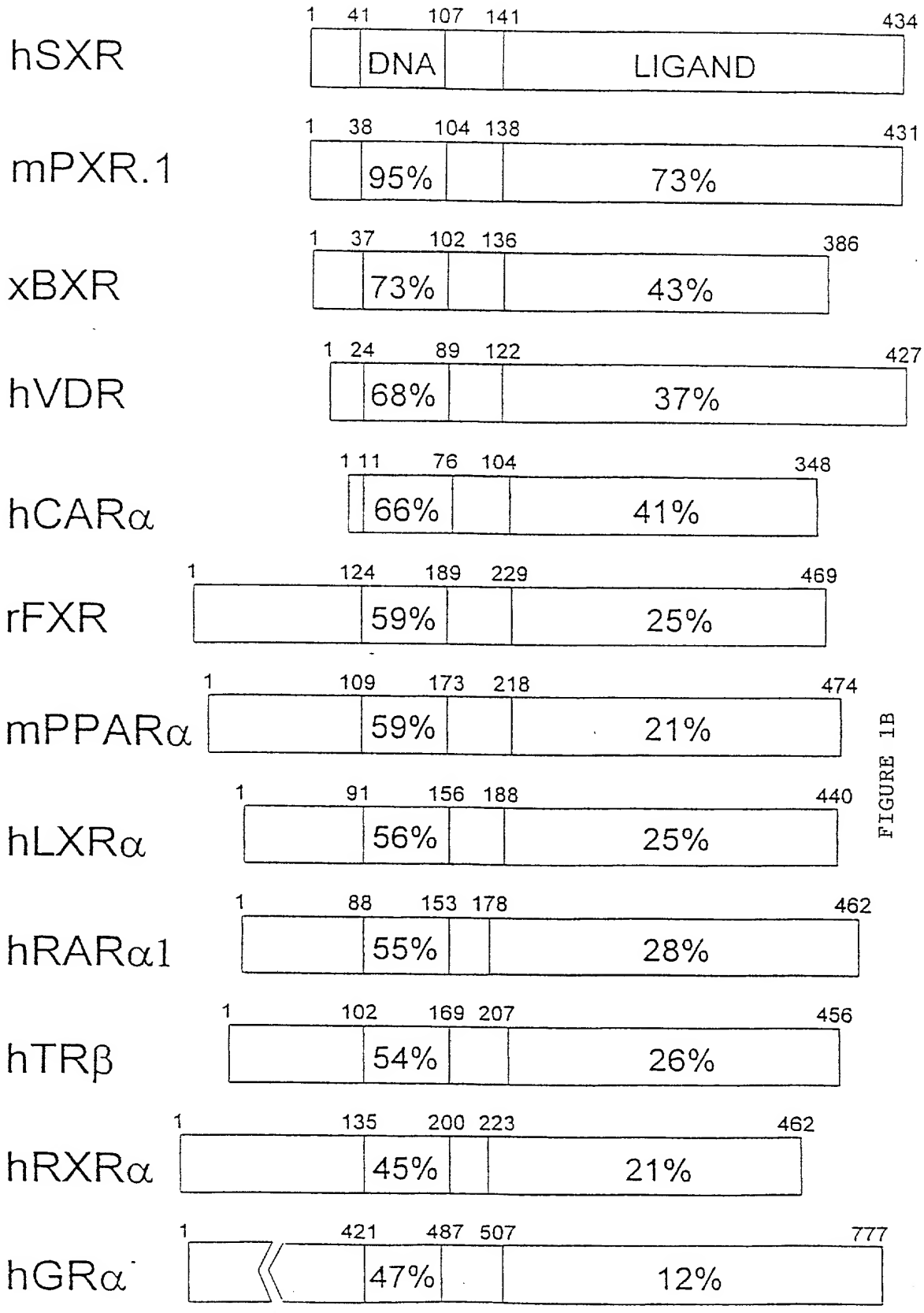

FIGURE 1B

1665455 - 09/05/06

200220 "SSST3001

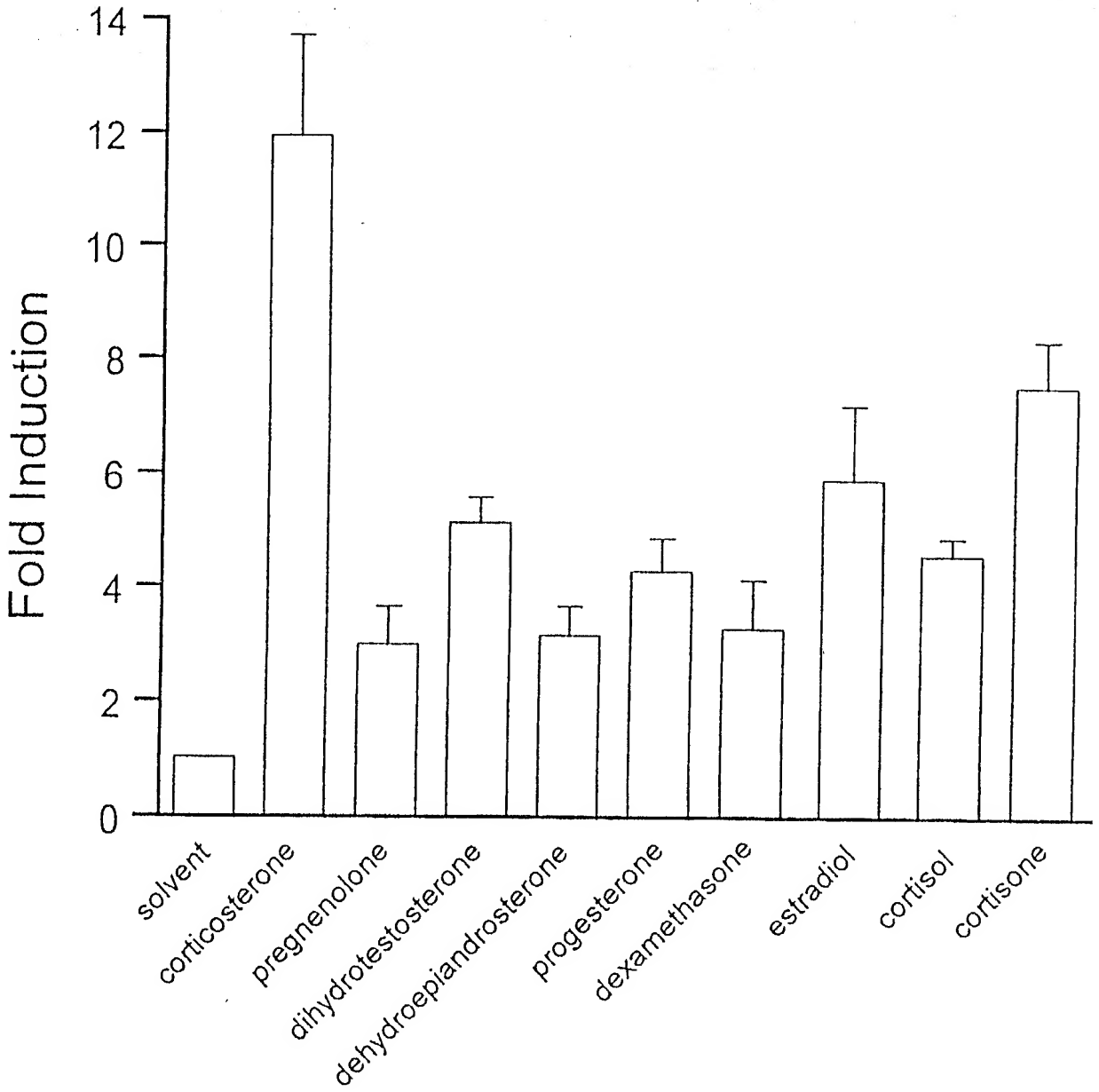

Figure 2

200220 SSST80T

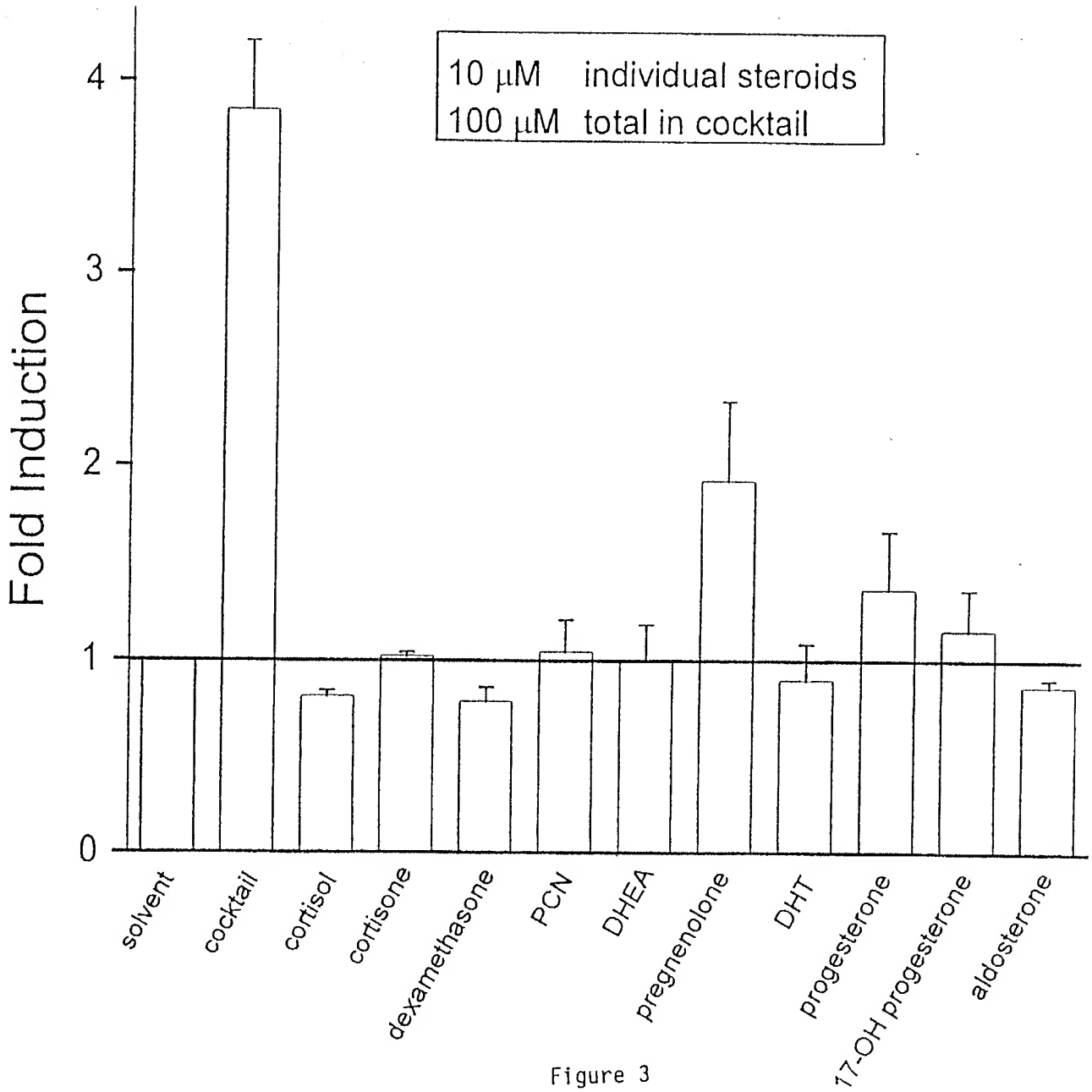

Figure 3

200220"5551800T

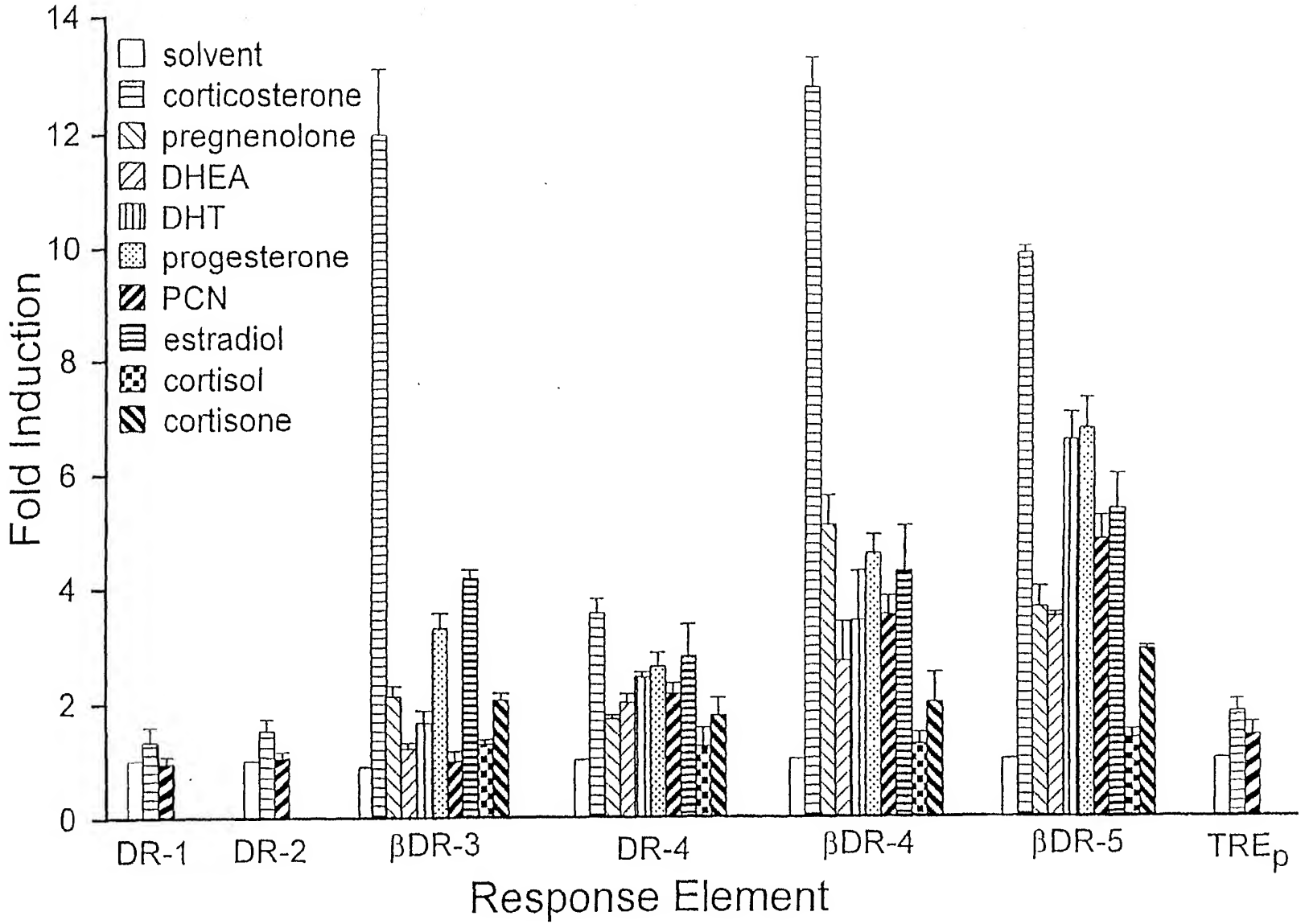

Figure 4

Applicant's Exhibit

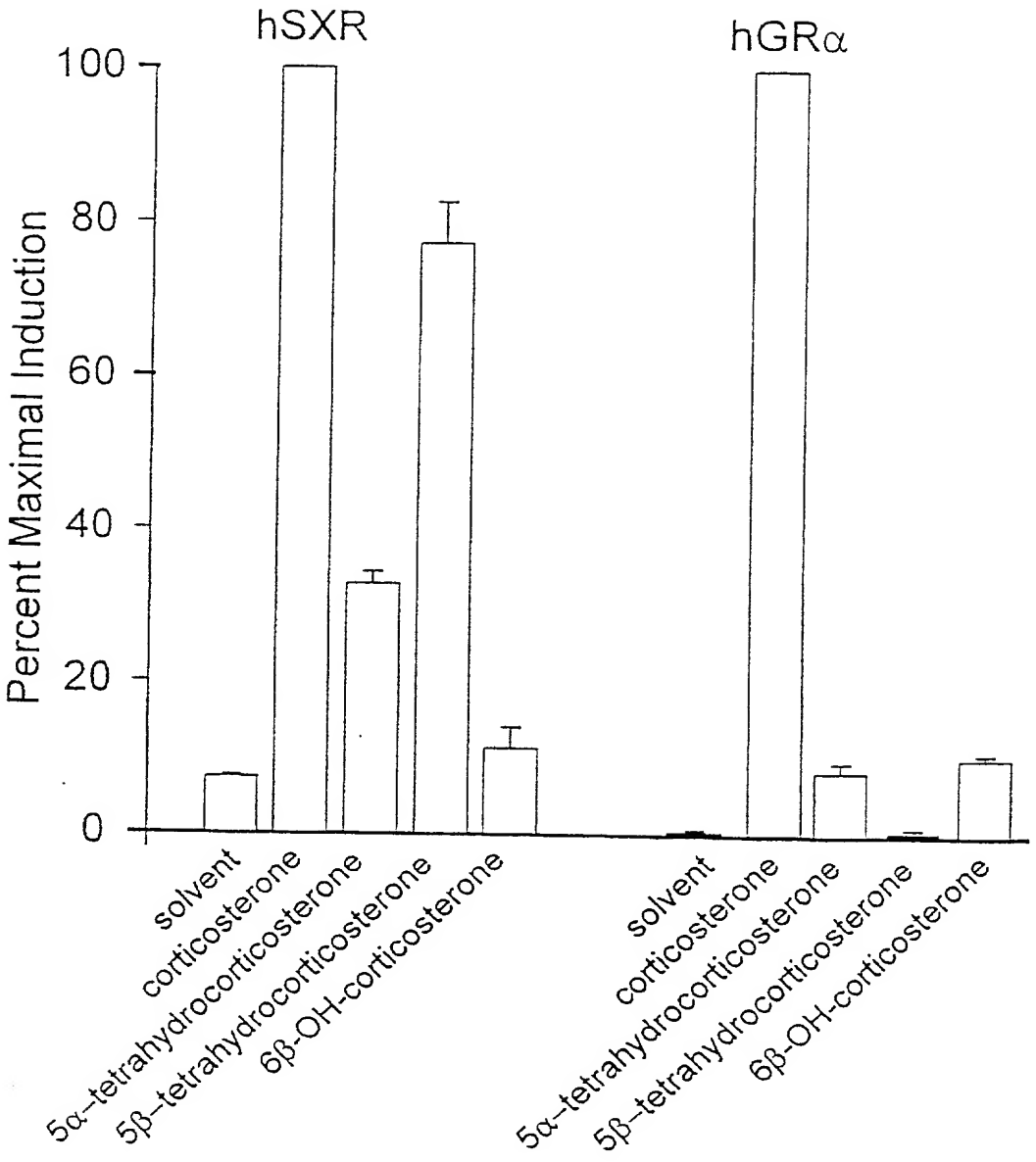

Figure 5.

300220 "SSSTBOOT

DR-3

rCYP3A1	tagac **AGTTCA** tga **AGTTCA** tctac
rCYP3A2	taagc **AGTTCA** taa **AGTTCA** tctac
rUGT1A6	actgt **AGTTCA** taa **AGTTCA** catgg

DR-4

rbCYP2C1	caatc **AGTTCA** acag **GGTTCA** ccaat
rP450R	cac **AGGTGA** gctg **AGGCCA** gcagc **AGGTCG** aaa

DR-5

rCYP2A1	gtgca **GGTTCA** actgg **AGGTCA** acatg
rCYP2A2	gtgct **GGTTCA** actgg **AGGTCA** gtatg
rCYP2C6	agtct **AGTTCA** gtggg **GGTTCA** gtctt
hCYP2E1	gagat **GGTTCA** aggaa **GGGTCA** ttaac

Figure 6A

CYP3A4 tagaata **TGAACT** caaagg **AGGTCA** gtgagtgg
CYP3A5 tagaata **TGAACT** caaagg **AGGTAA** gcaaaggg
CYP3A7 tagaata **TTAACT** caatgg **AGGC.A** gtgagtgg

Figure 6B

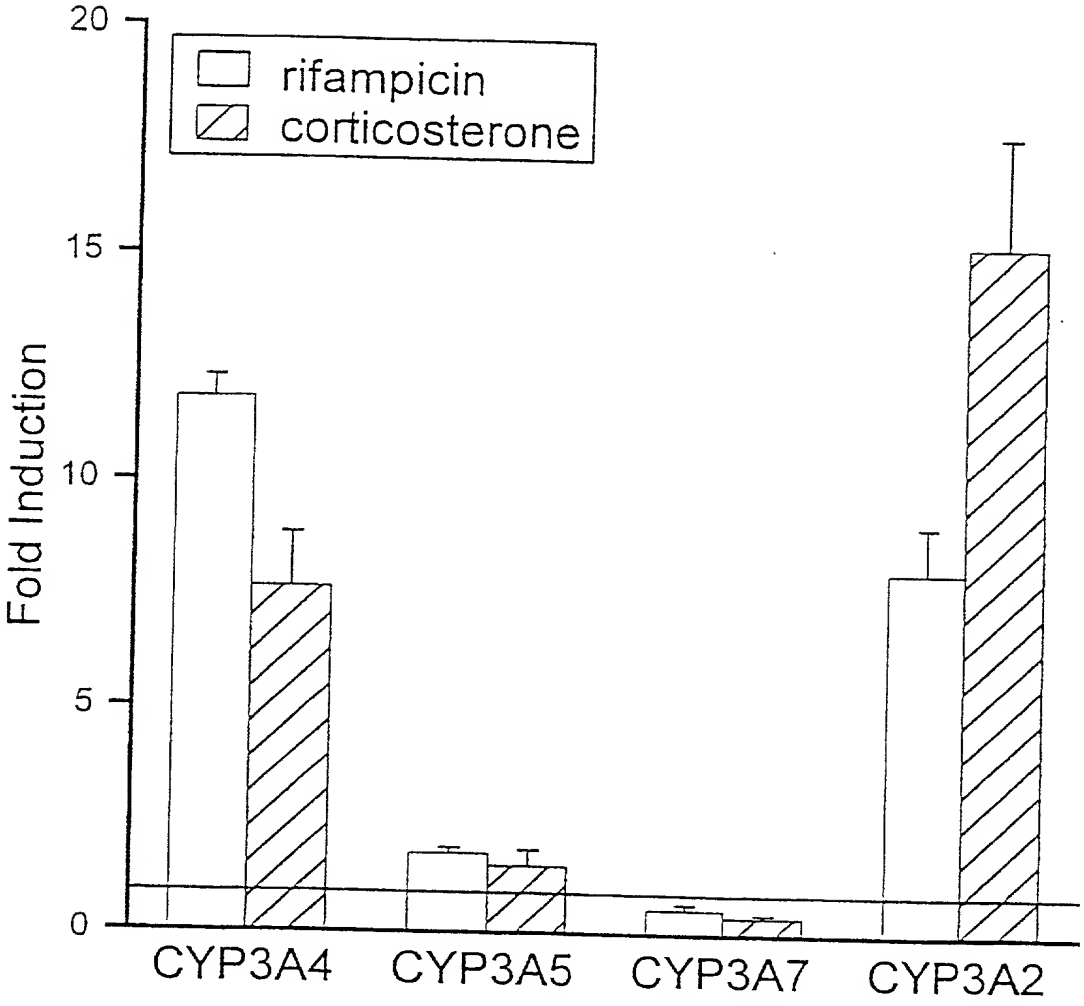
20
15
10
5
0
Fold Induction
rifampicin
corticosterone
CYP3A4
CYP3A5
CYP3A7
CYP3A2

Figure 6C

20020055504

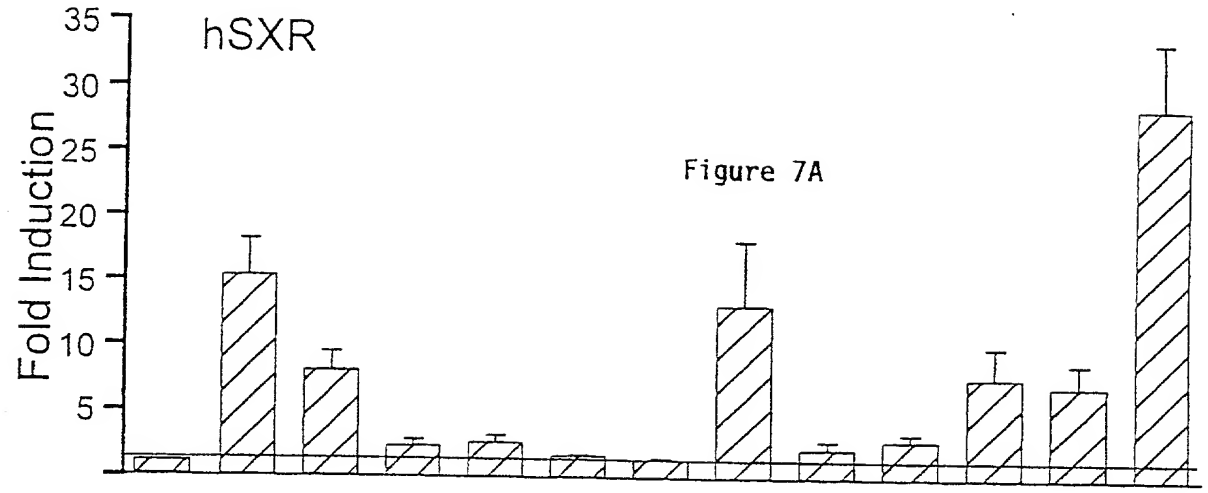

Figure 7A

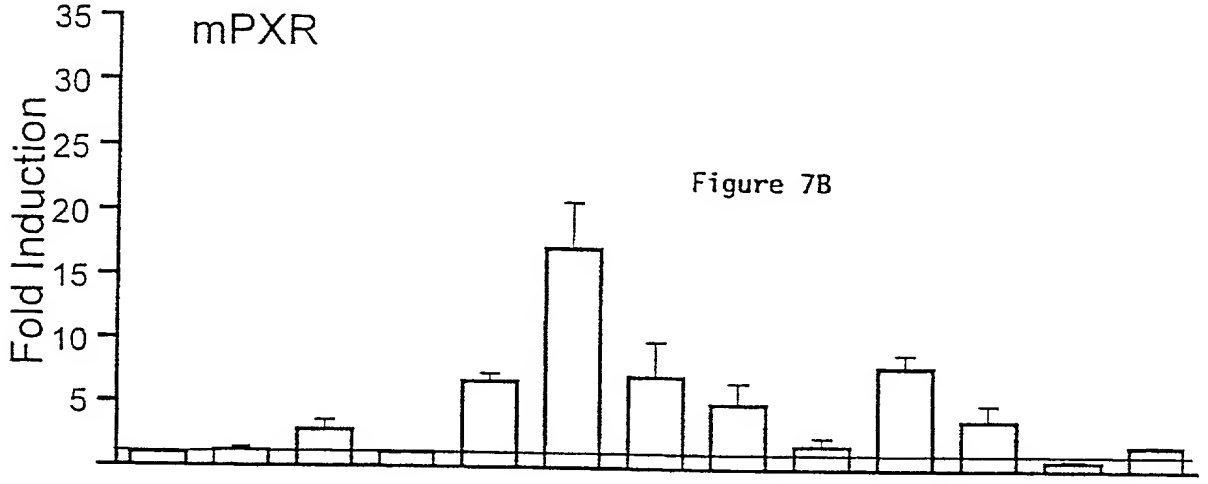

Figure 7B

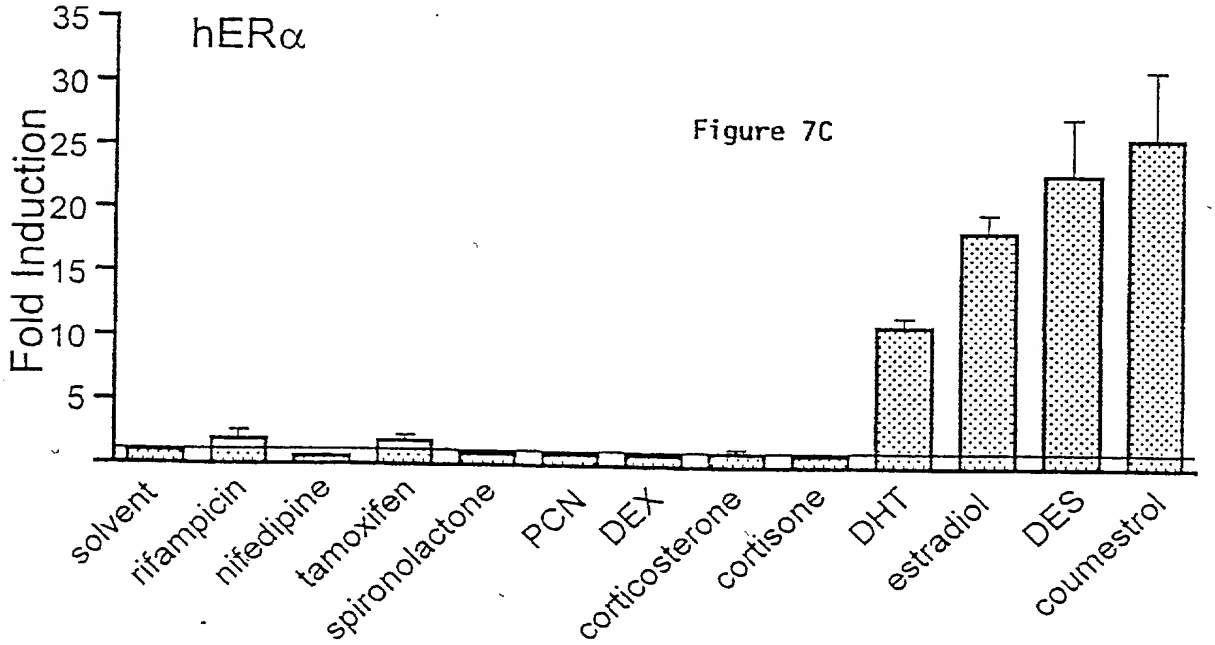

Figure 7C

20020-55407

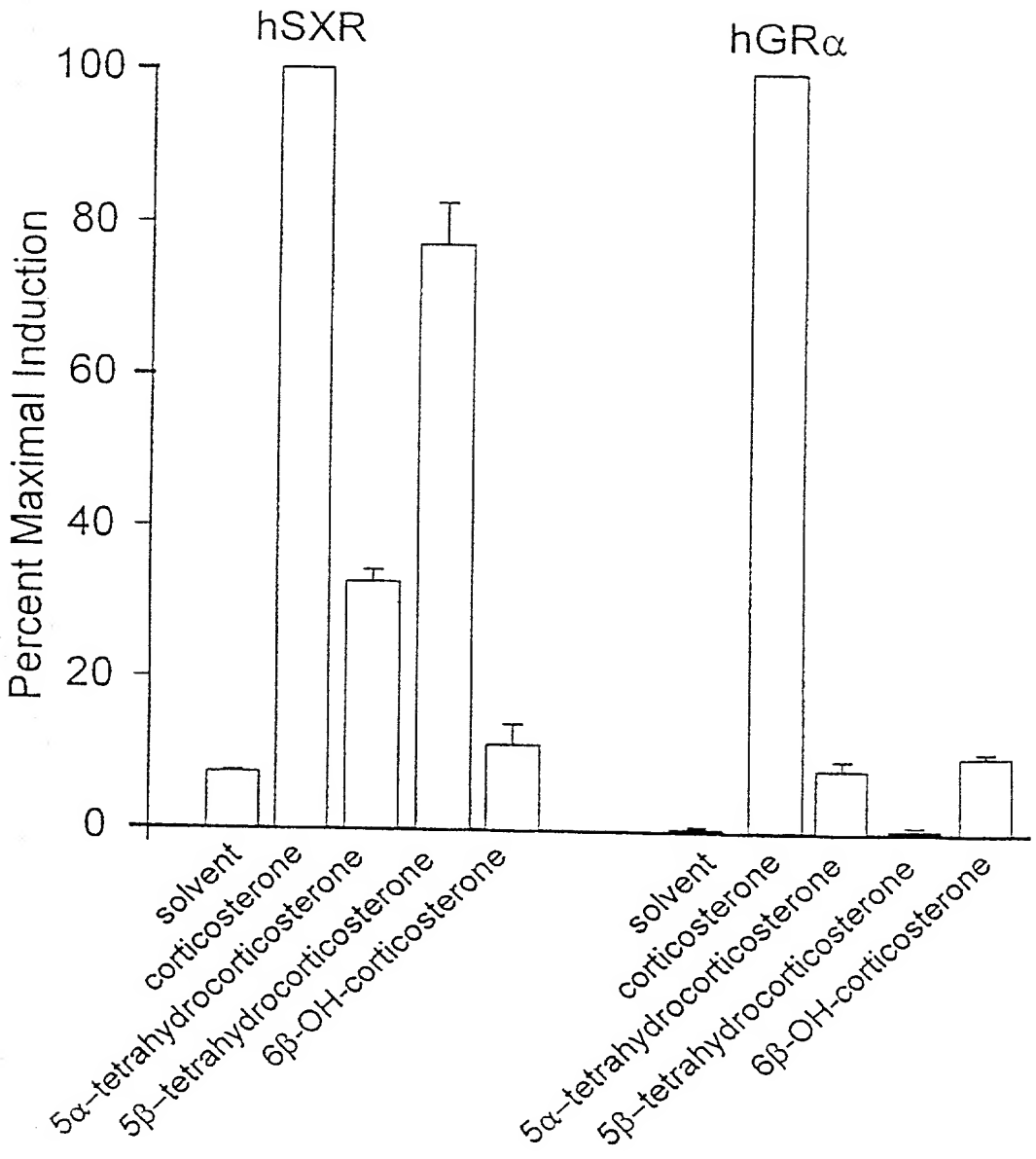

Figure 7D

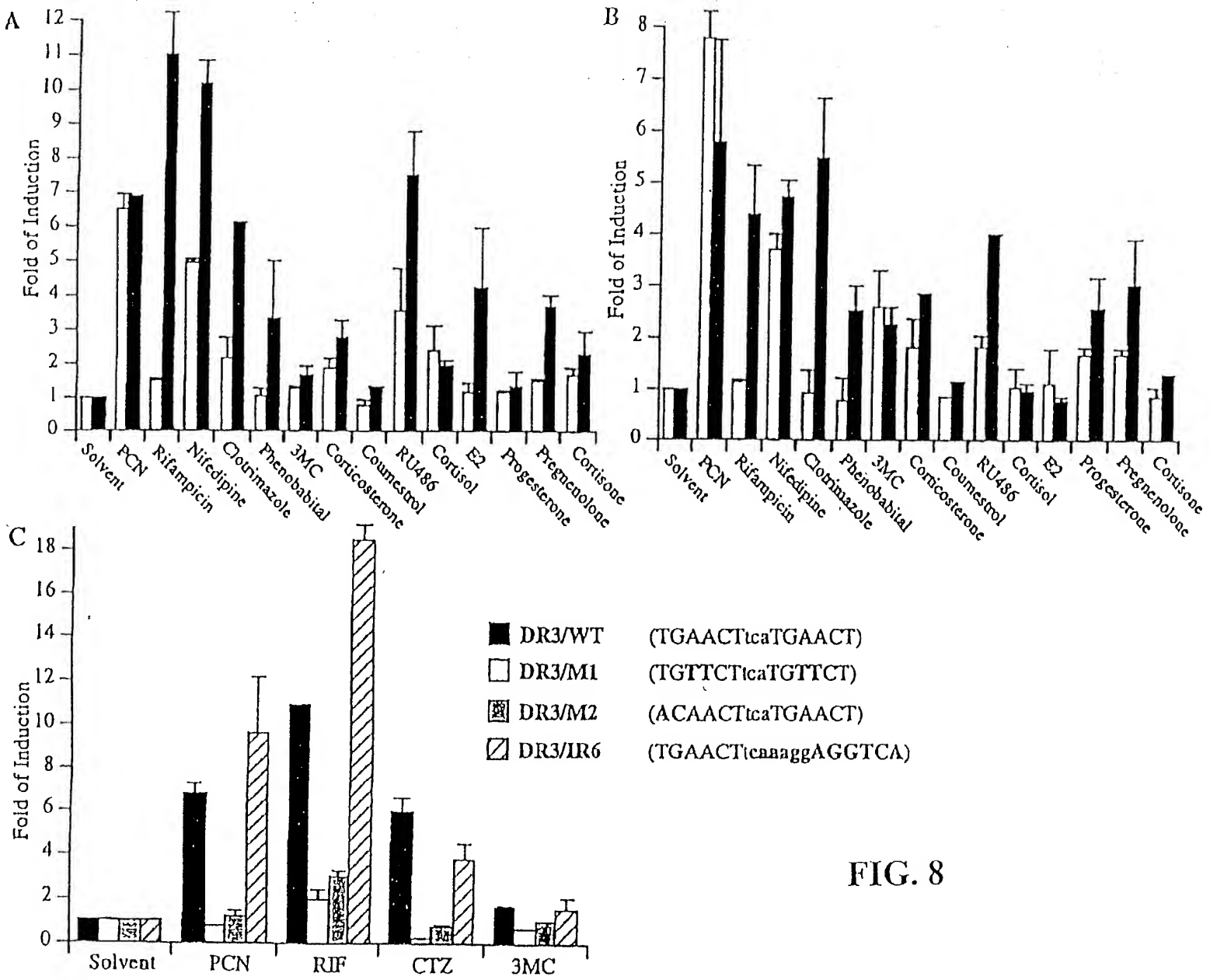
A
Fold of Induction
Solvent
PCN
Rifampicin
Nifedipine
Clotrimazole
Phenobabital
3MC
Corticosterone
Coumestrol
RU486
Cortisol
E2
Progesterone
Pregnenolone
Cortisone
B
Fold of Induction
Solvent
PCN
Rifampicin
Nifedipine
Clotrimazole
Phenobabital
3MC
Corticosterone
Coumestrol
RU486
Cortisol
E2
Progesterone
Pregnenolone
Cortisone
C
Fold of Induction
Solvent
PCN
RIF
CTZ
3MC
DR3/WT (TGAACTtcaTGAACT)
DR3/M1 (TGTTCTtcaTGTTCT)
DR3/M2 (ACAACTtcaTGAACT)
DR3/IR6 (TGAACTtcaaaggAGGTCA)

FIG. 8

EP 1 554 401 B1

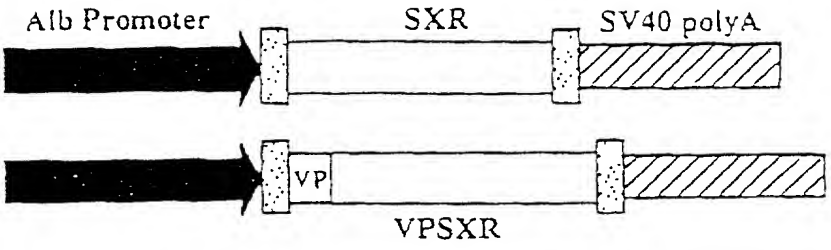

FIG. 9

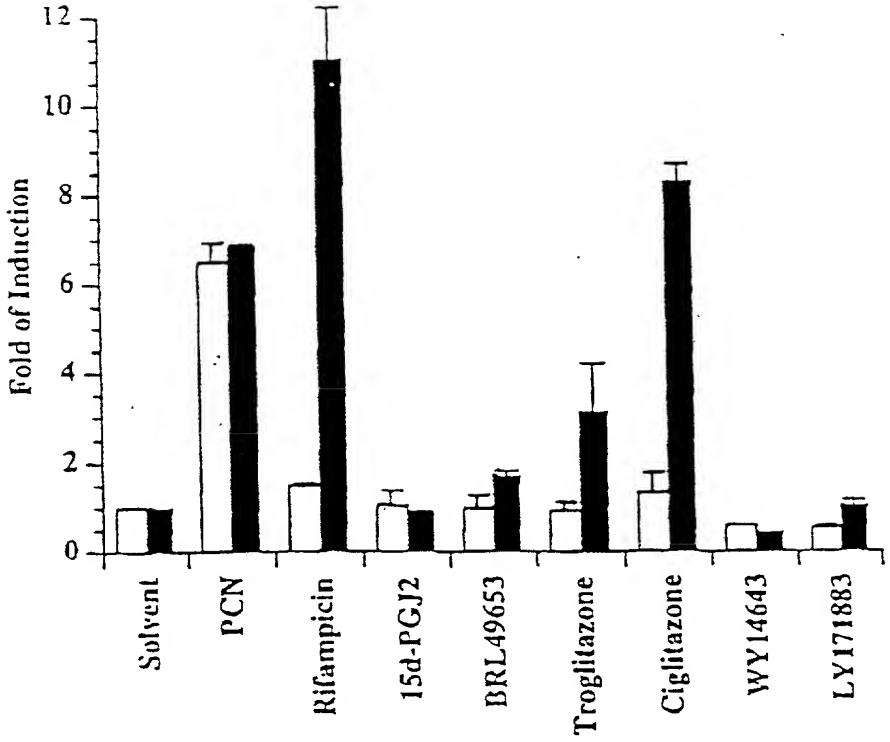

FIG. 10

1000241555.000205

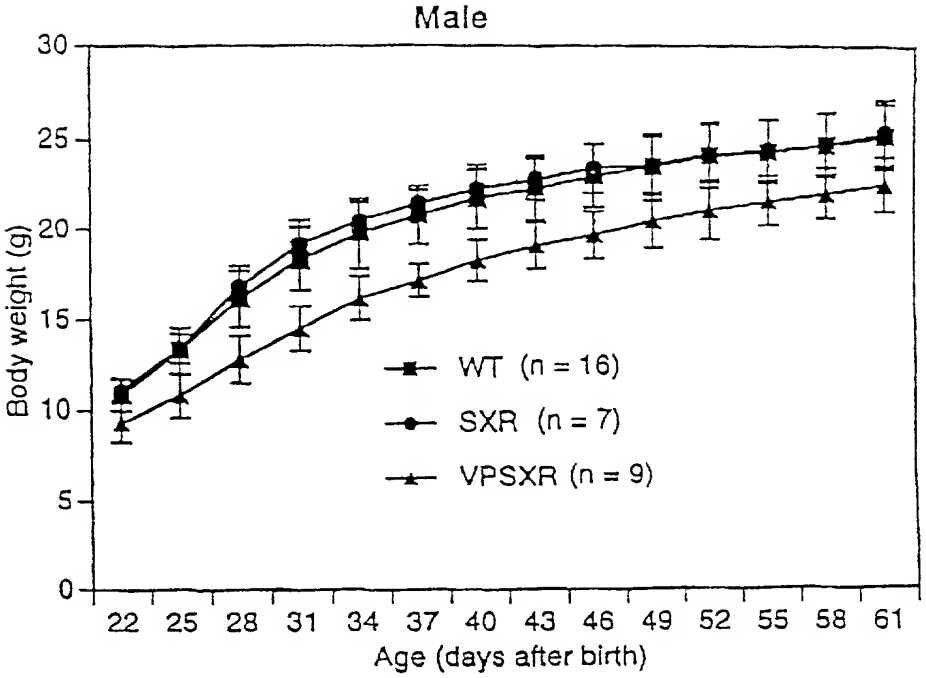

FIG. 11

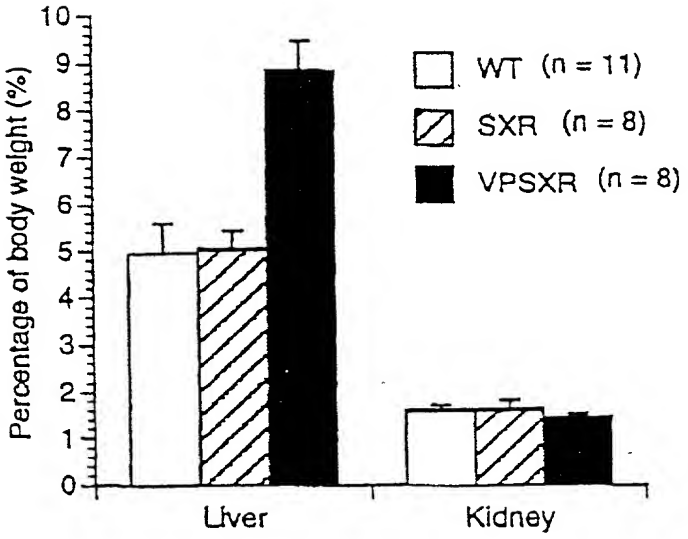

FIG. 12